

## INNOVATION IN GI MEDICINE

March 19, 2020

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This presentation contains forward-looking statements. All statements other than statements of historical facts contained in this presentation, including statements regarding our future results of operations and financial position, business strategy, prospective products, product approvals, research and development costs, timing and likelihood of success, plans and objectives of management for future operations, and future results of current and anticipated products, are forward-looking statements.

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This presentation also contains estimates and other statistical data made by independent parties and by us relating to market size and growth and other data about our industry. This data involves a number of assumptions and limitations, and you are cautioned not to give undue weight to such estimates. In addition, projections, assumptions, and estimates of our future performance and the future performance of the markets in which we operate are necessarily subject to a high degree of uncertainty and risk.

#### **Executive leadership team**



Tachi Yamada, MD Chairman

CMO & CSO, Takeda President Global Health, Gates Foundation Chairman R&D, GSK President, AGA



**Terrie Curran**CEO

President I&I Franchise, Celgene Led OTEZLA business from US launch through \$13b sale SVP Global Women's Health, Merck



Azmi Nabulsi, MD

Deputy CMO & CSO, Takeda Global Head Development, Takeda Division VP. Abbott



Martin Gilligan

CVP Marketing, Market Access, BD, Celgene I&I Franchise Executive Director Marketing, J&J Global Brand Leader, Merck



Joe Hand CAO

EVP, Global HR and Corporate Service, Celgene Executive Committee, Celgene Attorney at Jones Day



**Tom Harris**SVP, Regulatory
& Quality

SVP/Head of Global Regulatory, Takeda VP US Regulatory, Takeda Humira Global Project Head, Abbott



Aditya Kohli, PhD CBO

Principal, Frazier
VP BD, Scout Bio
Engagement Manager,
McKinsey



Eckhard Leifke, MD
CMO

CMO, Omeros Global Head CVM Early Project & External Opportunities, Sanofi



Larry Miller GC

General Counsel, Cyclerion SVP and General Counsel, Blue Buffalo Chief Counsel, Pfizer Consumer Healthcare



**David Socks** CFO

Venture Partner, Frazier
CEO, Outpost Medicine
COO, Incline Therapeutics
SVP, Cadence
Pharmaceuticals

#### POTENTIAL BREAKTHROUGH

for acid-related disorders

#### **VONOPRAZAN**



- > Potassium competitive acid blocker (P-CAB)
- > Potentially first-in-class in US, Europe, and Canada
- > US/EU/Canada rights licensed from Takeda
- > 18 Phase 3 studies completed by Takeda in >6,000 subjects
- Approved in 13 countries across Asia and Latin America
- > >\$500M net sales in Japan in fourth full year on the market



#### P-CABs: next generation of acid-control therapeutics





**Takecab**° (vonoprazan)

#### **P-CABs**

Introduced in 2015 in Japan

- > 24hr time above pH 4: >70%
- > >\$500M net sales in Japan

#### **PPIs** Introduced in 1989

- > 24hr time above pH 4: 40-71%
- > ~\$12.5b peak US sales

#### **ANTACIDS** Introduced in 1930s

> Short duration of efficacy

> 24hr time above pH 4: 33%

> ~\$3.5b peak US sales

#### **PPIs:** mechanism limits effectiveness

## **GASTRIC PARIETAL CELL** Secretory canaliculus Inactive phase Active phase after meal Proton pump proton pump



### PPI: COVALENTLY BINDING PRODRUG

Acid needed for activation but unstable in presence of acid

Meal required to stimulate pumps

Short plasma half-life of 1 to 2 hours

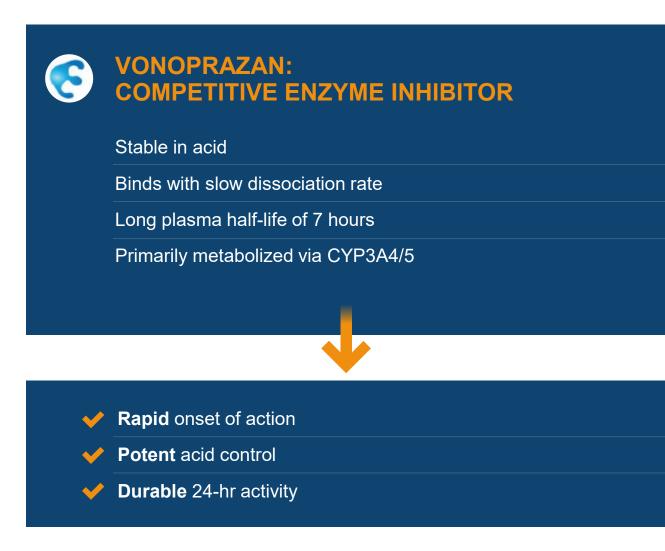
Primarily metabolized via CYP2C19



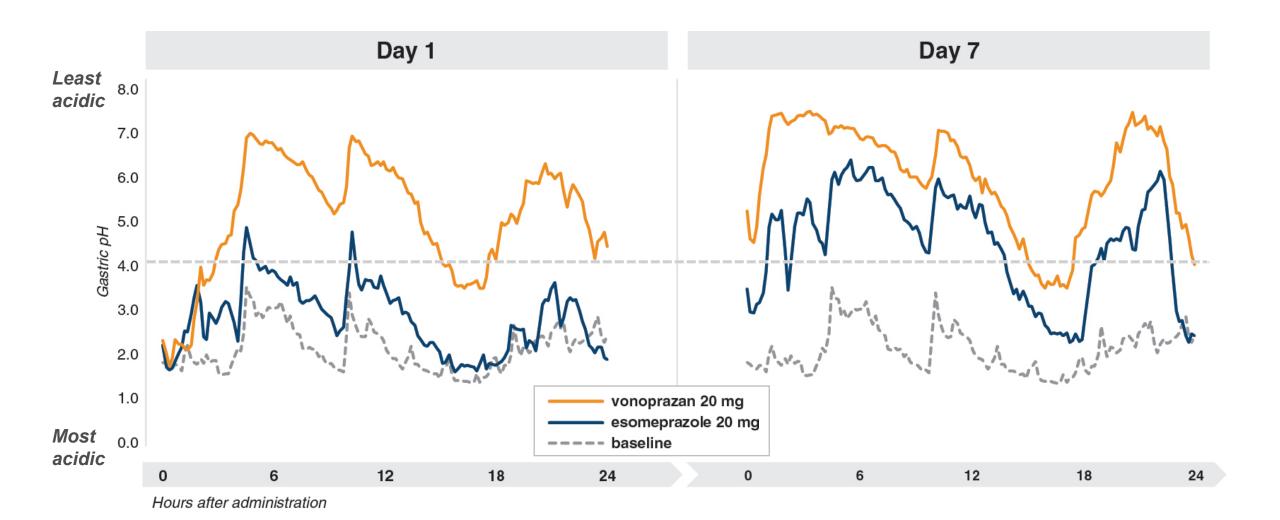
- X Slow onset of action
- Limited potency
- Limited duration of activity

#### Vonoprazan: distinct mechanism designed to address PPI shortcomings

## **GASTRIC PARIETAL CELL** Secretory canaliculus Inactive phase Active phase after meal Proton pump

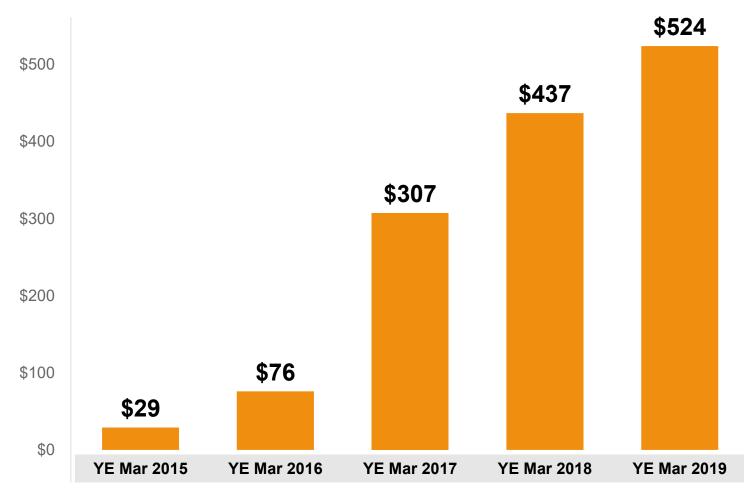


#### Vonoprazan demonstrated faster and more potent acid control vs. PPI in healthy volunteers



## Vonoprazan achieved **RAPID ADOPTION** and strong sales growth in Japan TAKECAB® (VONOPRAZAN) JAPAN LAUNCH **FEBRUARY 2015** Takecab® is a registered trademark of Takeda Pharmaceutical Co. Ltd.

#### **VONOPRAZAN NET SALES, US\$ MILLION\***

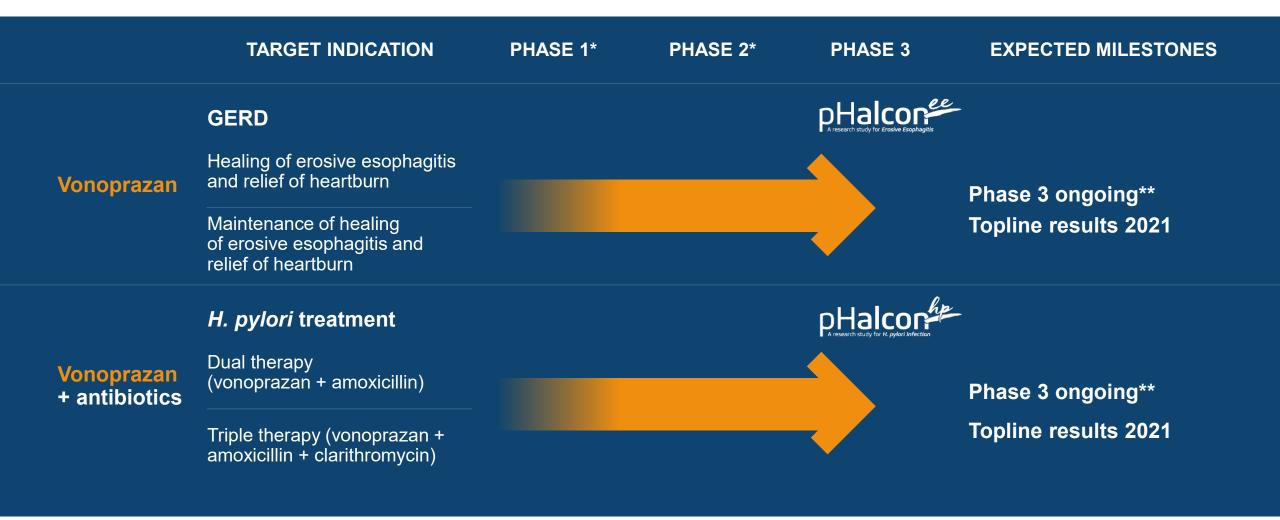


Note: vonoprazan net sales of approximately \$337M for the six months ended December 31, 2019

\*U.S. dollars based on the June 30, 2019 conversion rate of 0.009 dollar to one yen



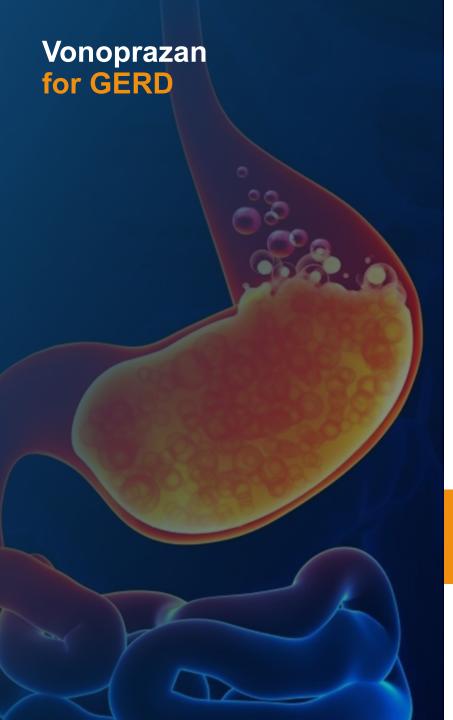
#### Phathom pipeline



Phathom has development and commercialization rights to vonoprazan in the United States, Europe, and Canada \*Phase 1 and 2 studies conducted by Takeda



<sup>\*\*</sup> On March 19, 2020, temporarily paused new patient randomization in view of COVID-19 global pandemic



~65M US and ~50M EU5 patients with GERD

6.1 billion PPI doses prescribed in US for the 12 months ended May 31, 2019

~15-45% inadequately treated with PPIs

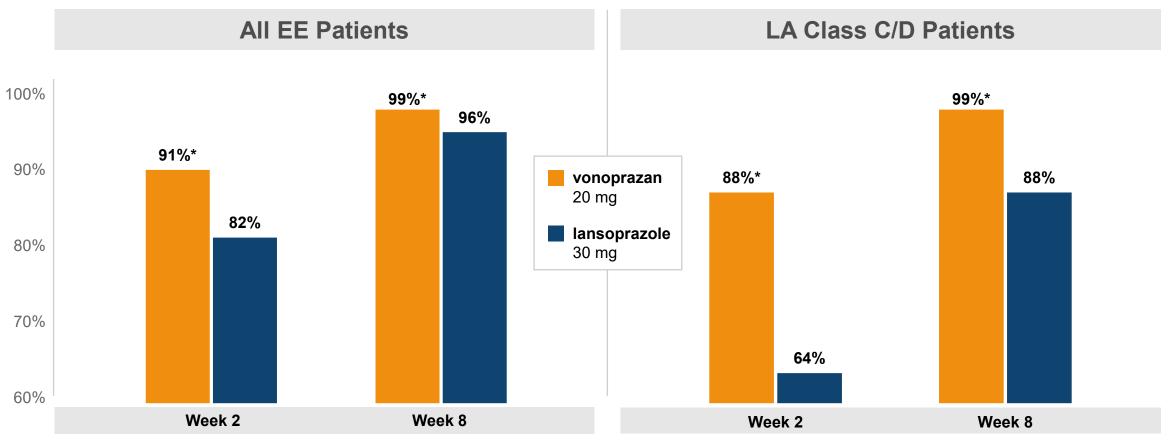
Many patients experience breakthrough heartburn and recurrence of erosions while on PPIs

Vonoprazan may offer more rapid, potent, and durable healing and symptom control

El-Serag APT 2010; El-Serag Gut 2014; IQVIA data July 2019

#### Japan erosive esophagitis (EE) Phase 3: demonstrated faster and improved healing vs. PPI

#### Patients with healed EE, %

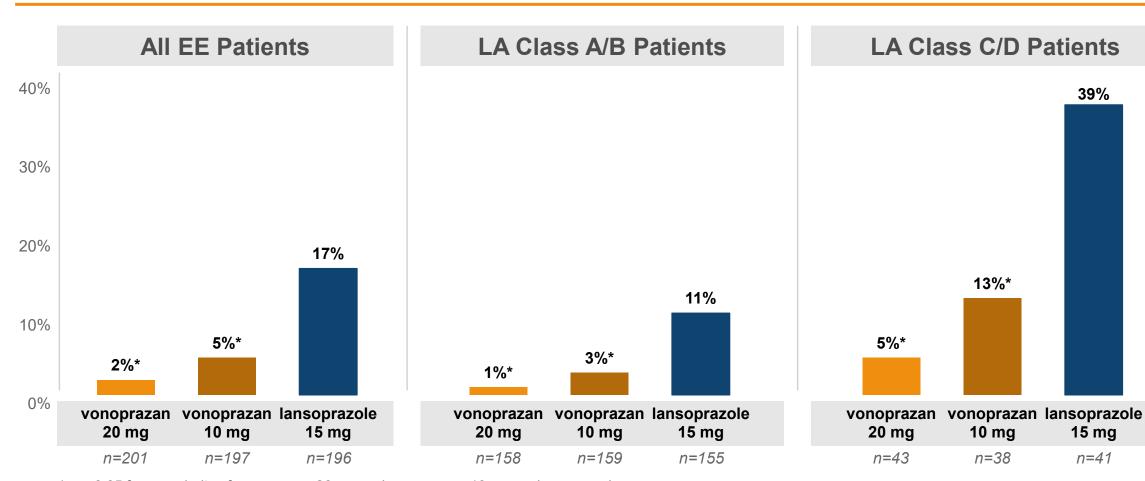


n=404, \*Statistically significant (p < 0.05) for superiority versus lansoprazole

n=147, \*Statistically significant (p < 0.05) for superiority versus lansoprazole

#### Japan erosive esophagitis (EE) Phase 3: demonstrated lower recurrence rates vs. PPI

#### Patients with recurrence of EE at 6 months, %



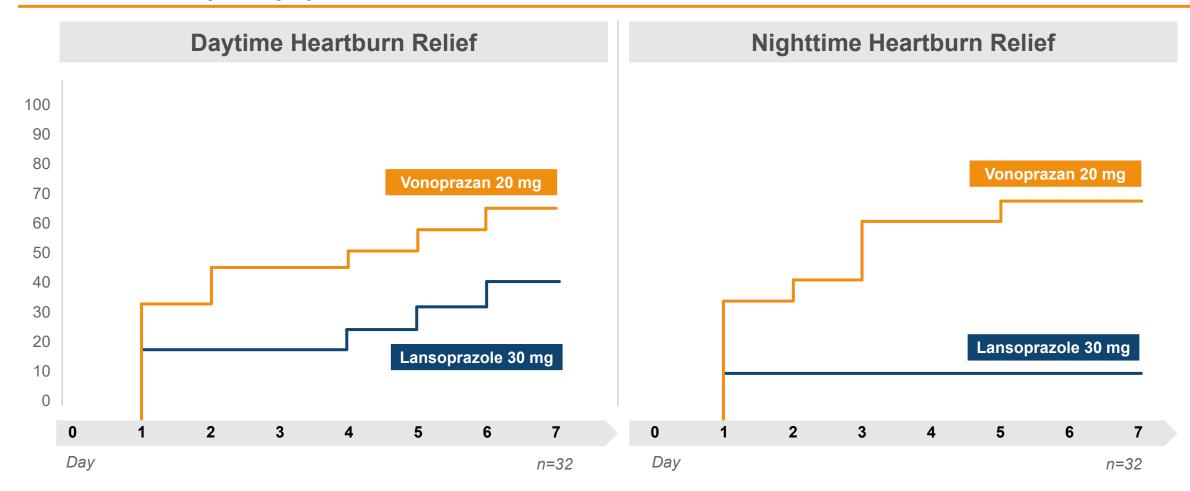
<sup>\*</sup> p < 0.05 for superiority of vonoprazan 20 mg and vonoprazan 10 mg vs. lansoprazole

n = 41

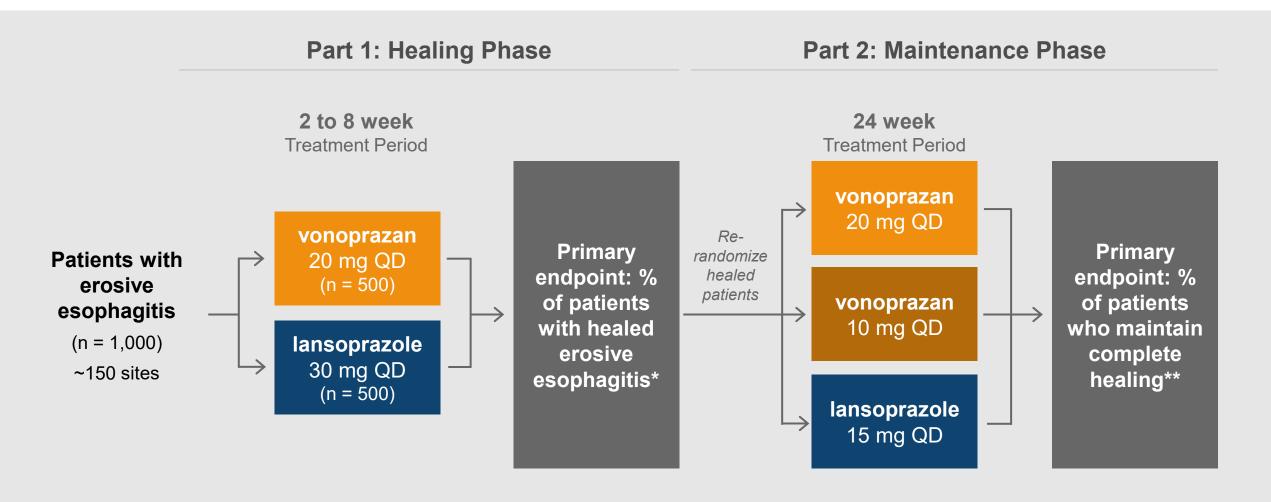
39%

#### Faster and more complete heartburn relief of vonoprazan vs. PPI

#### Patients with complete symptom relief, %



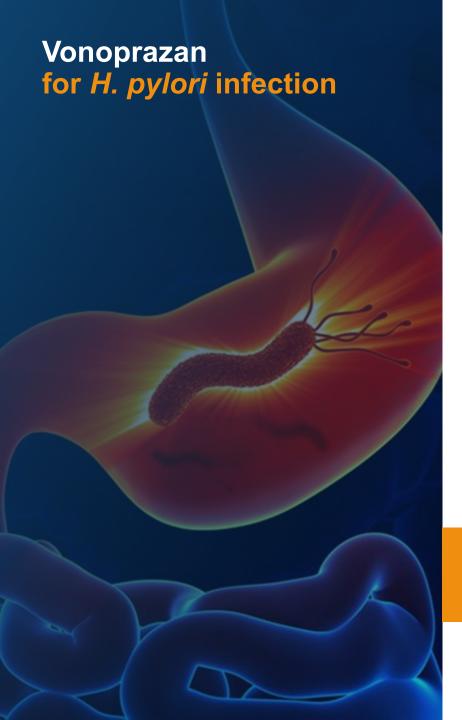
#### Phathom US/Europe EE Phase 3 study design



Note: Diagnosis and healing of erosive esophagitis confirmed by endoscopy

<sup>\*</sup>Primary analysis of non-inferiority; key secondary analysis assessing superiority at week 2 in Los Angeles class C/D patients

<sup>\*\*</sup>Primary analysis of non-inferiority; if non-inferiority met, superiority will also be assessed



~115M US and ~145M EU5 patients with *H. pylori* 

~2.5M US patients treated annually

H. pylori designated as a Class I carcinogen by WHO and Qualifying Pathogen under FDA GAIN Act

Eradication rates have fallen to <80% due to increasing antibiotic resistance

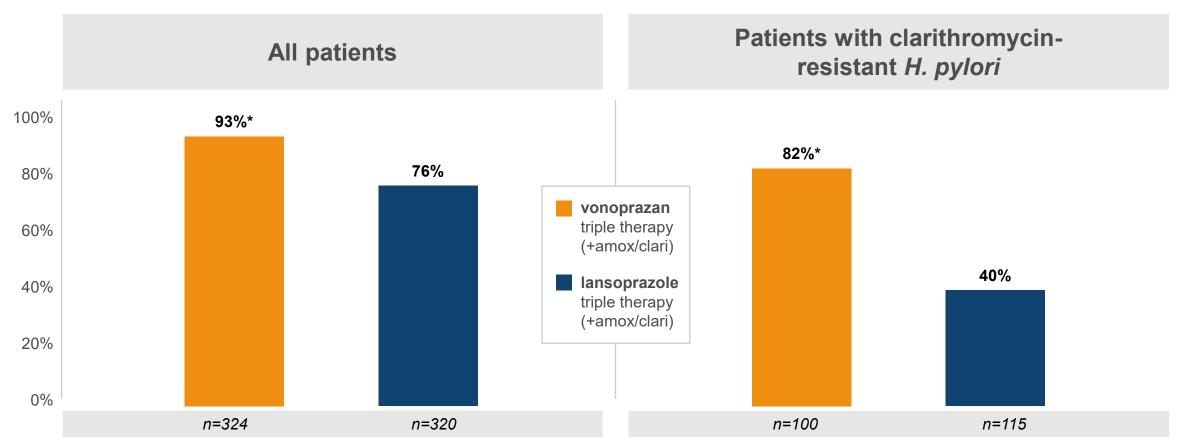
Antibiotic potency increases at higher pH

Vonoprazan-based regimens may restore eradication rates above 90% in the US and Europe

Hooi Gastroenterology 2017; Graham et al 2018; Erah et al 1997

#### Japan H. pylori Phase 3: vonoprazan triple therapy demonstrated superiority to PPI therapy

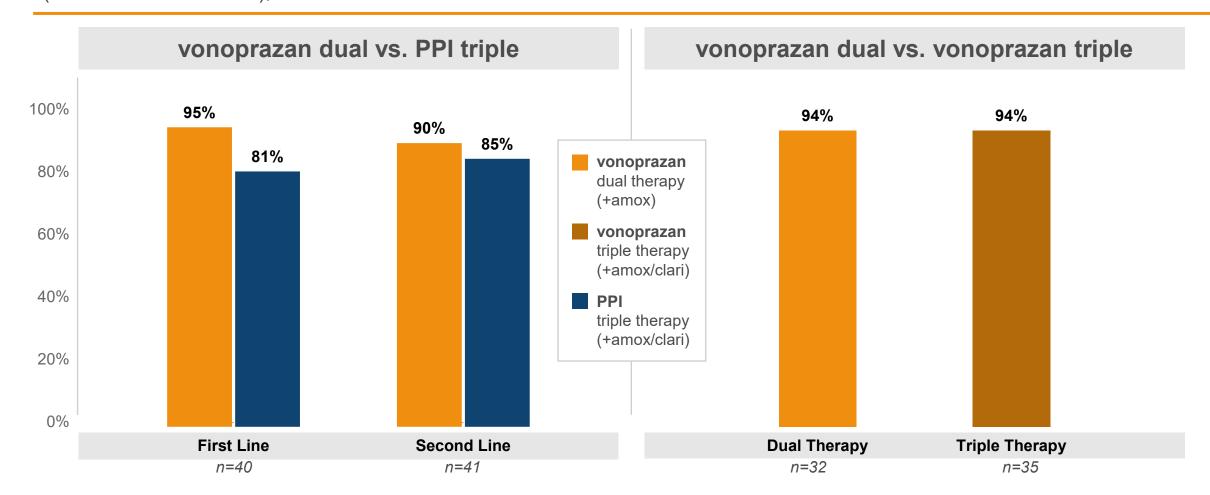
### First-line triple therapy eradication rates of *H. pylori* (combo with amoxicillin/clarithromycin), %



<sup>\*</sup> p < 0.0001 for superiority of vonoprazan-based triple therapy to lansoprazole-based triple therapy

#### Vonoprazan dual therapy also demonstrated >90% H. pylori eradication

Eradication rates of *H. pylori* (dual or triple therapy) (combo with antibiotics), %

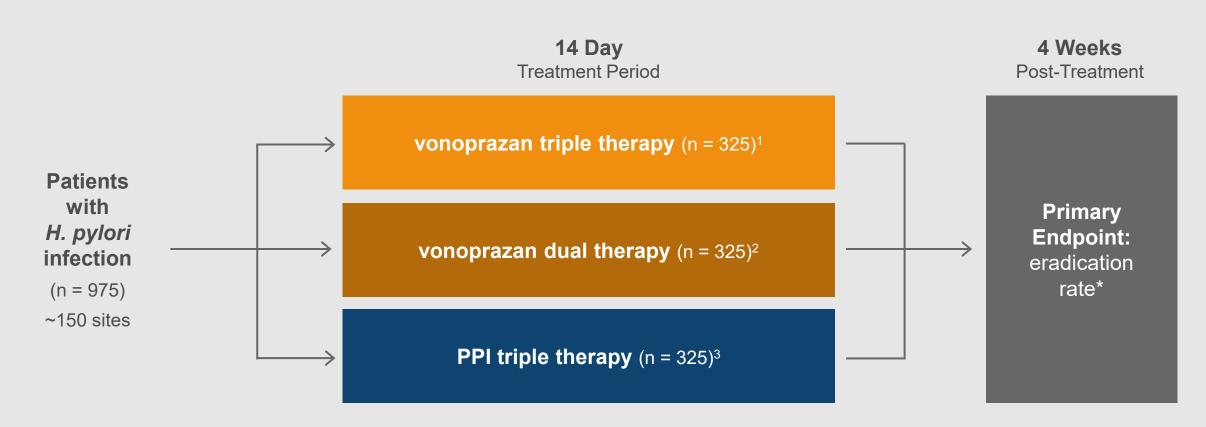


H. pylori eradication rates in Japan have increased since the launch of vonoprazan

VONOPRAZAN-BASED REGIMENS ACHIEVED ~80% SHARE IN JAPAN BY 2016



#### Phathom US/Europe H. pylori Phase 3 study design



<sup>1.</sup> vonoprazan 20 mg BID + amoxicillin 1 g BID + clarithromycin 500 mg BID

Note: Diagnosis of infection and test of cure confirmed by <sup>13</sup>C-urea breath test

<sup>2.</sup> vonoprazan 20 mg BID + amoxicillin 1 g TID (partially blinded)

<sup>3.</sup> lansoprazole 30 mg BID + amoxicillin 1 g BID + clarithromycin 500 mg BID

<sup>\*</sup>Primary analysis of non-inferiority excluding patients with infection resistant to clarithromycin and amoxicillin; key secondary analyses of superiority in patients with clarithromycin resistant infection and in all comers

# Vonoprazan safety profile SIMILAR TO PPIS

**6,683 subjects** received vonoprazan in clinical studies

No dose-related increase in adverse events observed

>25 million patients received vonoprazan since launch

### ADVERSE EVENTS IN CLINICAL DEVELOPMENT REFLECTED IN JAPANESE PRESCRIBING INFORMATION

Incidence of 0.1-5.0%

Diarrhea <sup>1</sup>	Elevated liver enzymes
Constipation	Rash
Nausea	Eosinophilia

### HEPATIC EVENTS OF SPECIFIC INTEREST IN LIGHT OF FIRST-GENERATION PCABs

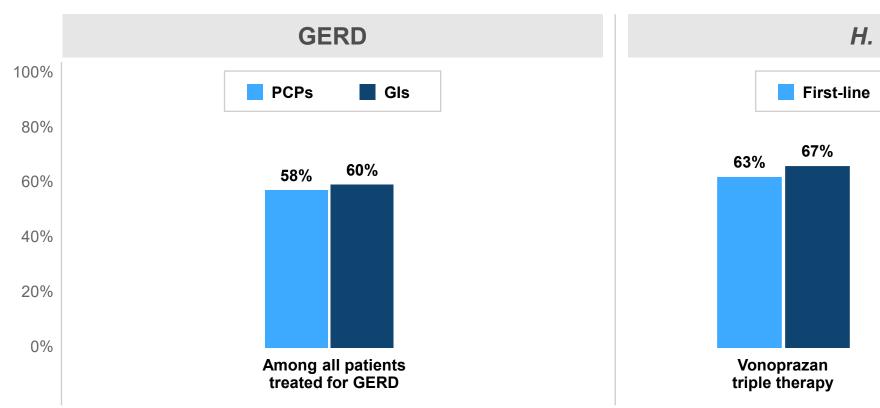
Pooled data across head-to-head Phase 2 and 3 studies	vonoprazan 10 and 20mg	lansoprazole 15 and 30mg
ALT or AST > 3X ULN or Bilirubin >2X ULN	1.0%	0.8%

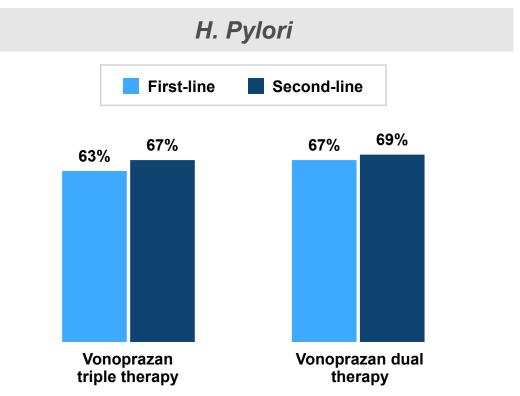
<sup>1. 10.6%</sup> in combination with antibiotics for treatment of *H. pylori* 



#### US physicians have strong preference to prescribe vonoprazan

#### US physician preference share, %

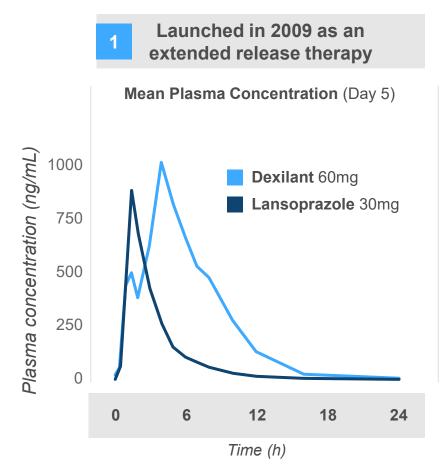


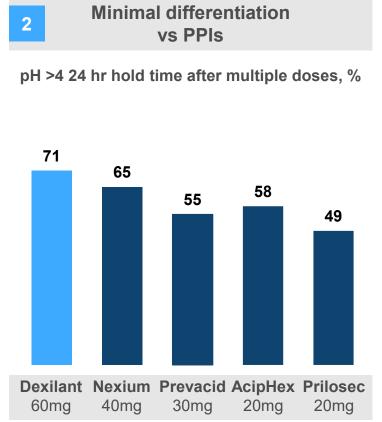


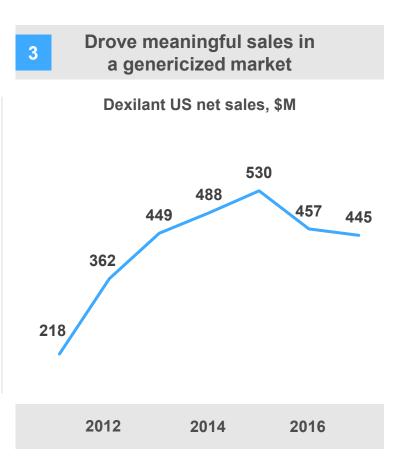
2019 US survey of 100 gastroenterologists and 100 primary care physicians

#### Dexilant case study: last of the branded PPIs









#### **Dexilant case study:** market access

\$9.42/dose US WAC<sup>1</sup>

~90% of commercial and ~80% of Medicare covered lives have access to Dexilant<sup>2</sup>

65% of commercial covered lives have unrestricted access without step edits or prior authorization<sup>2</sup>

35% of commercial covered lives have access at the lowest branded cost tier<sup>2</sup>

## FORMULARY STATUS AMONG TOP 5 PLANS By covered lives<sup>3</sup>

HEALTH PLAN	COVERAGE
Aetna Self-Insured	Tier 2 Preferred
Cigna Standard 3-Tier (National)	Tier 2 Preferred
CVS Caremark Advanced Control Specialty	Tier 2 Preferred
Express Scripts National Preferred	Tier 3 Non-Preferred
UnitedHealthcare Advantage 3-Tier	Tier 3 Non-Preferred

NO STEP-EDITS OR PRIOR AUTHORIZATION

<sup>2</sup> MMIT formulary lookup tool as of June 25, 2019

<sup>3</sup> Fingertip Formulary Accessed 4Q18

#### Financial highlights

Cash and Cash Equivalents (as of 12/31/2019) <sup>1</sup>	\$243.8M
Debt (as of 12/31/2019) <sup>1,2</sup>	\$25.0M
Common Shares Issued (as of 3/12/2020) <sup>1</sup>	28,964,506

<sup>&</sup>lt;sup>1</sup> December 31, 2019 Form 10-K

<sup>&</sup>lt;sup>2</sup> Silicon Valley Bank Term Loan. Increased to \$50.0M on March 16, 2020 drawdown of additional \$25M

# Phathom. PHARMACEUTICALS

#### **NASDAQ: PHAT**

- ✓ Significant unmet medical need
- Established safety and efficacy in Japan
- ✓ Late-stage US/EU program
- Large commercial opportunity
- Seasoned team and investors